

SEQUENCE LISTING

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CENTRE NATIONAL DE LA RECHERCHE SCIENTIFIQUE

<120> GRB14 AND THE INSULIN RECEPTOR AND SCREENING OF NOVEL MEDICINES

<130> 45636-5051

<140> US 09/936,697

<141> 2001-09-17

<160> 28

<170> PatentIn Ver. 2.1

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<212> PRT

<213> Rattus sp.

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Gly Gln Lys Thr Arg Val Ile Asp Asn Pro Thr Glu Ala Leu Ser Val \$20\$

Ala Val Glu Glu Gly Leu Ala Trp Arg Lys Lys

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Val Ser Glu Asn Ser Leu Val Ala Met Asp Phe Ser Gly Gln Lys Thr \$20\$

Arg Val Ile Asp Asn Pro Thr Glu Ala Leu Ser Val Ala Val Glu Glu 35 \$40\$

Gly Leu Ala Trp Arg Lys Lys Gly Cys Leu Arg Leu Gly Asn His Gly 50 55 60

Ser Pro Thr Ala Pro Ser Gln Ser Ser Ala Val Asn Met Ala Leu His 65 70 75 80

Arg Ser Gln Pro

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Ser Pro Thr Ala Pro Ser Gln Ser Ser Ala Val Asn Met Ala Leu His
Arg Ser Gln Pro Trp Phe His His Arg Ile Ser Arg Asp Glu Ala Gln
Gln Leu Ile Thr Arg Gln Gly Pro Val Asp Gly Val Phe Leu Val Arg
Asp Ser Gln Ser Asn Pro Arg Thr Phe Val Leu Ser Met Ser His Gly
                            120
Gln Lys Ile Lys His Phe Gln Ile Ile Pro Val Glu Asp Asp Gly Glu
                        135
Val Phe His Thr Leu Asp Asp Gly His Thr Lys Phe Thr Asp Leu Ile
Gln Leu Val Glu Phe Tyr Gln Leu Asn Lys Gly Val Leu Pro Cys Lys
Leu Lys His Tyr Cys Ala Arg Met Ala Val
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Gly Gln Lys Ser Arg Val Ile Glu Asn Pro Thr Glu Ala Leu Ser Val
Ala Val Glu Glu Gly Leu Ala Trp Arg Lys Lys
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Arg Val Ile Glu Asn Pro Thr Glu Ala Leu Ser Val Ala Val Glu Glu
Gly Leu Ala Trp Arg Lys Lys Gly Cys Leu Arg Leu Gly Thr His Gly
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Ser Pro Thr Ala Ser Ser Gln Ser Ser Ala Thr Asn Met Ala Ile His 70 Arg Ser Gln Pro <210 > 7 <211> 174 <212> PRT <213> Homo sapiens <400> 7 Pro Met Arg Ser Ile Ser Glu Asn Ser Leu Val Ala Met Asp Phe Ser Gly Gln Lys Ser Arg Val Ile Glu Asn Pro Thr Glu Ala Leu Ser Val Ala Val Glu Glu Gly Leu Ala Trp Arg Lys Lys Gly Cys Leu Arg Leu Gly Thr His Gly Ser Pro Thr Ala Ser Ser Gln Ser Ser Ala Thr Asn Met Ala Ile His Arg Ser Gln Pro Trp Phe His His Lys Ile Ser Arg Asp Glu Ala Gln Arg Leu Ile Ile Gln Gln Gly Leu Val Asp Gly Val Phe Leu Val Arg Asp Ser Gln Ser Asn Pro Lys Thr Phe Val Leu Ser 105 Met Ser His Gly Gln Lys Ile Lys His Phe Gln Ile Ile Pro Val Glu Asp Asp Gly Glu Met Phe His Thr Leu Asp Asp Gly His Thr Arg Phe 135 Thr Asp Leu Ile Gln Leu Val Glu Phe Tyr Gln Leu Asn Lys Gly Val 160 145 Leu Pro Cys Lys Leu Lys His Tyr Cys Ala Arg Ile Ala Leu 170 <210> 8 <211> 186 <212> PRT

<213> Homo sapiens

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Ile Ser Glu Asn Ser Leu Val Ala Met Asp Phe Ser Gly Gln Lys Ser 25 20

Arg Val Tle Glu Asn Pro Thr Glu Ala Leu Ser Val Ala Val Glu Glu Gly Leu Ala Trp Arg Lys Lys Gly Cys Leu Arg Leu Gly Thr His Gly Ser Pro Thr Ala Ser Ser Gln Ser Ser Ala Thr Asn Met Ala Ile His Arg Ser Gln Pro Trp Phe His His Lys Ile Ser Arg Asp Glu Ala Gln Arg Leu Ile Ile Gln Gln Gly Leu Val Asp Gly Val Phe Leu Val Arg Asp Ser Gln Ser Asn Pro Lys Thr Phe Val Leu Ser Met Ser His Gly 120 Gln Lvs Ile Lvs His Phe Gln Ile Ile Pro Val Glu Asp Asp Gly Glu 135 Met Phe His Thr Leu Asp Asp Gly His Thr Arg Phe Thr Asp Leu Ile 145 150 Gln Leu Val Glu Phe Tyr Gln Leu Asn Lys Gly Val Leu Pro Cys Lys Leu Lys His Tyr Cys Ala Arg Ile Ala Leu <210> 9 <211> 43 <212> PRT <213> mus muris <400> 9 Pro Met Arg Ser Val Ser Glu Asn Ser Leu Val Ala Met Asp Phe Ser Gly Gln Ile Gly Arg Val Ile Asp Asn Pro Ala Glu Ala Gln Ser Ala Ala Leu Glu Glu Gly His Ala Trp Arg Asn Gly 40 <210> 10 <211> 82 <212> PRT <213> mus muris Pro Gln Arg Lys Gly Leu Pro Pro Pro Phe Asn Ala Pro Met Arg Ser Val Ser Glu Asn Ser Leu Val Ala Met Asp Phe Ser Gly Gln Ile Gly

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Arg Val Tle Asp Asn Pro Ala Glu Ala Gln Ser Ala Ala Leu Glu Glu 35 40 45

Gly His Ala Trp Arg Asn Gly Ser Thr Arg Met Asn Ile Leu Ser Ser 50 55 60

Gln Ser Pro Leu His Pro Ser Thr Leu Asn Ala Val Ile His Arg Thr 65 70 75 80

Gln His

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<211> 172 <212> PRT

<213> mus muris

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Pro Met Arg Ser Val Ser Glu Asn Ser Leu Val Ala Met Asp Phe Ser 1 5 10 15

Gly Gln Ile Gly Arg Val Ile Asp Asn Pro Ala Glu Ala Gln Ser Ala 20 25 30

Ala Leu Glu Glu Gly His Ala Trp Arg Asn Gly Ser Thr Arg Met Asn 35 40 45

Ile Leu Ser Ser Gln Ser Pro Leu His Pro Ser Thr Leu Asn Ala Val 50 55 60

Ile His Arg Thr Gln His Trp Phe His Gly Arg Ile Ser Arg Glu Glu 65 70 75 80

Ser His Arg Ile Ile Lys Gln Gln Gly Leu Val Asp Gly Leu Phe Leu 85 90 95

Leu Arg Asp Ser Gln Ser Asn Pro Lys Ala Phe Val Leu Thr Leu Cys 100 105 \cdot 110

His His Gln Lys Ile Lys Asn Phe Gln Ile Leu Pro Cys Glu Asp Asp 115 120 125

Gly Gln Thr Phe Phe Thr Leu Asp Asp Gly Asn Thr Lys Phe Ser Asp 130 135 140

Leu Ile Gln Leu Val Asp Phe Tyr Gln Leu Asn Lys Gly Val Leu Pro 145 150 155 160

Cys Lys Leu Lys His His Cys Ile Arg Val Ala Leu 165 170

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<212> PRT

<213> mus muris

<400> 12

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Val Ser Glu Asn Ser Leu Val Ala Met Asp Phe Ser Gly Gln Ile Gly
Arg Val Ile Asp Asn Pro Ala Glu Ala Gln Ser Ala Ala Leu Glu Glu
Gly His Ala Trp Arg Asn Gly Ser Thr Arg Met Asn Ile Leu Ser Ser
Gln Ser Pro Leu His Pro Ser Thr Leu Asn Ala Val Ile His Arg Thr
Gln His Trp Phe His Gly Arg Ile Ser Arg Glu Glu Ser His Arg Ile
Ile Lys Gln Gln Gly Leu Val Asp Gly Leu Phe Leu Leu Arg Asp Ser
                                105
Gln Ser Asn Pro Lys Ala Phe Val Leu Thr Leu Cys His His Gln Lys
                            120
Ile Lys Asn Phe Gln Ile Leu Pro Cys Glu Asp Asp Gly Gln Thr Phe
                        135
Phe Thr Leu Asp Asp Gly Asn Thr Lys Phe Ser Asp Leu Ile Gln Leu
                    150
Val Asp Phe Tyr Gln Leu Asn Lys Gly Val Leu Pro Cys Lys Leu Lys
His His Cys Ile Arg Val Ala Leu
            180
<210> 13
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<213> Homo sapiens
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Gly Gln Thr Gly Arg Val Ile Glu Asn Pro Ala Glu Ala Gln Ser Ala
Ala Leu Glu Glu Gly His Ala Trp Arg Lys Arg
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<210> 16 <211> 184 <212> PRT <213> Homo sapiens <400> 16 Gln Gln Arg Lys Ala Leu Leu Ser Pro Phe Ser Thr Pro Val Arg Ser Val Ser Glu Asn Ser Leu Val Ala Met Asp Phe Ser Gly Gln Thr Gly Arg Val Ile Glu Asn Pro Ala Glu Ala Gln Ser Ala Ala Leu Glu Glu Gly His Ala Trp Arg Lys Arg Ser Thr Arg Met Asn Ile Leu Gly Ser Gln Ser Pro Leu His Pro Ser Thr Leu Ser Thr Val Ile His Arg Thr Gln His Trp Phe His Gly Arg Phe Ser Arg Glu Glu Ser His Arg Ile Ile Lys Gln Gln Gly Leu Val Asp Gly Leu Phe Leu Leu Arg Asp Ser 105 Gln Ser Asn Pro Lys Ala Phe Val Leu Thr Leu Cys His His Gln Lys 120 115 Ile Lys Asn Phe Gln Ile Leu Pro Cys Glu Asp Asp Gly Gln Thr Phe 135 Phe Ser Leu Asp Asp Gly Asn Thr Lys Phe Ser Asp Leu Ile Gln Leu 145 Val Asp Phe Tyr Gln Leu Asn Lys Gly Val Leu Pro Cys Lys Leu Lys 170 His His Cys Ile Arg Val Ala Leu 180 <210> 17 <211> 43 <212> PRT <213> Rattus sp. <400> 17 Pro Leu Arg Ser Val Ser Asp Asn Thr Leu Val Ala Met Asp Phe Ser 10 Gly His Ala Gly Arg Val Ile Asp Asn Pro Arg Glu Ala Leu Ser Ala Ala Met Glu Glu Ala Gln Ala Trp Arg Lys Lys

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150

Leu Arg His Cys Cys Thr Arg Val Ala Leu 165 170

<210> 24

<211> 182

<212> PRT <213> Homo sapiens

<400> 24

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Ala Ser Asp Asn Thr Leu Val Ala Met Asp Phe Ser Gly His Ala Gly \$20\$

Arg Val Ile Glu Asn Pro Arg Glu Ala Leu Ser Val Ala Leu Glu Glu 35 40 45

Ala Gln Ala Trp Arg Lys Lys Thr Asn His Arg Leu Ser Leu Pro Met 50 60

Pro Ala Ser Gly Thr Ser Leu Ser Ala Ala Ile His Arg Thr Gln Leu 65 75 80

Trp Phe His Gly Arg Ile Ser Arg Glu Glu Ser Gln Arg Leu Ile Gly 85 90 95

Gln Gln Gly Leu Val Asp Gly Leu Phe Leu Val Arg Glu Ser Gln Arg 100 105 110

Asn Pro Gln Gly Phe Val Leu Ser Leu Cys His Leu Gln Lys Val Lys 115 120 125

His Tyr Leu Ile Leu Pro Ser Glu Glu Glu Gly Arg Leu Tyr Phe Ser 130 140

Met Asp Asp Gly Gln Thr Arg Phe Thr Asp Leu Leu Gln Leu Val Glu 145 150 155 160

Phe His Gln Leu Asn Arg Gly Ile Leu Pro Cys Leu Leu Arg His Cys \$165\$ 170 175

Cys Thr Arg Val Ala Leu 180

<210> 25

<211> 43 <212> PRT

<213> mus muris

<400> 25

Pro Leu Arg Ser Val Ser Asp Asn Thr Leu Val Ala Met Asp Phe Ser

Gly His Ala Gly Arg Val Ile Asp Asn Pro Arg Glu Ala Leu Ser Ala 20 25 30 Ala Met Glu Glu Ala Gln Ala Trp Arg Lys Lys 35 40

<210> 26

<211> 80 <212> PRT

<213> mus muris

<400> 26

Ser Arg His Leu Arg Leu Ser Tyr Leu Gly Ser Pro Pro Leu Arg Ser 1 $$ 15

Val Ser Asp Asn Thr Leu Val Ala Met Asp Phe Ser Gly His Ala Gly \$20\$

Arg Val Ile Asp Asn Pro Arg Glu Ala Leu Ser Ala Ala Met Glu Glu 35 40 45

Ala Gln Ala Trp Arg Lys Lys Thr Asn His Arg Leu Ser Leu Pro Thr 50 55 60

Thr Cys Ser Gly Ser Ser Leu Ser Ala Ala Ile His Arg Thr Gln Pro 65 70 75 80

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<210> 27

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<213> mus muris

<400> 27

Pro Leu Arg Ser Val Ser Asp Asn Thr Leu Val Ala Met Asp Phe Ser 1 10 15

Gly His Ala Gly Arg Val Ile Asp Asn Pro Arg Glu Ala Leu Ser Ala 20 25 30

Ala Met Glu Glu Ala Gln Ala Trp Arg Lys Lys Thr Asn His Arg Leu 35 40 45

Ser Leu Pro Thr Thr Cys Ser Gly Ser Ser Leu Ser Ala Ala Ile His 50 60

Arg Thr Gln Pro Trp Phe His Gly Arg Ile Ser Arg Glu Glu Ser Gln 65 70 75 80

Arg Leu Ile Gly Gln Gln Gly Leu Val Asp Gly Val Phe Leu Val Arg 85 90 95

Glu Ser Gln Arg Asn Pro Gln Gly Phe Val Leu Ser Leu Cys His Leu 100 105 110

Gln Lys Val Lys His Tyr Leu Ile Leu Pro Ser Glu Asp Glu Gly Cys 115 120 125

Leu Tyr Phe Ser Met Asp Glu Gly Gln Thr Arg Phe Thr Asp Leu Leu 130 \$135\$

Gln Leu Val Glu Phe His Gln Leu Asn Arg Gly Ile Leu Pro Cys Leu 155 150 Leu Arg His Cys Cys Ala Arg Val Ala Leu 165 <210> 28 <211> 182 <212> PRT <213> mus muris <400> 28 Ser Arg His Leu Arg Leu Ser Tyr Leu Gly Ser Pro Pro Leu Arg Ser Val Ser Asp Asn Thr Leu Val Ala Met Asp Phe Ser Gly His Ala Gly Arg Val Ile Asp Asn Pro Arg Glu Ala Leu Ser Ala Ala Met Glu Glu Ala Gln Ala Trp Arg Lys Lys Thr Asn His Arg Leu Ser Leu Pro Thr Thr Cys Ser Gly Ser Ser Leu Ser Ala Ala Ile His Arg Thr Gln Pro Trp Phe His Gly Arg Ile Ser Arg Glu Glu Ser Gln Arg Leu Ile Gly Gln Gln Gly Leu Val Asp Gly Val Phe Leu Val Arg Glu Ser Gln Arg Asn Pro Gln Gly Phe Val Leu Ser Leu Cys His Leu Gln Lys Val Lys 120 His Tyr Leu Ile Leu Pro Ser Glu Asp Glu Gly Cys Leu Tyr Phe Ser Met Asp Glu Gly Gln Thr Arg Phe Thr Asp Leu Leu Gln Leu Val Glu Phe His Gln Leu Asn Arg Gly Ile Leu Pro Cys Leu Leu Arg His Cys 170 Cys Ala Arq Val Ala Leu

180